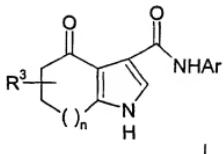


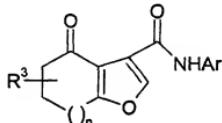
We claim:

1. A method of preparing a compound of the formula:



I

comprising reacting a compound of the formula:



II

5 with an excess of ammonia source in a reaction inert solvent at an elevated temperature until reaction is complete;

wherein Ar is phenyl or heterocycle, said phenyl or heterocycle being substituted with -O-(CH₂)_m-NR¹R², -O(CH₂)C(O)OR⁴, -CH(NR⁵R⁶)CH₃, or OH, and said phenyl or heterocycle being optionally substituted with one or two groups selected from C₁ - C₆ alkoxy, C₁ - C₆ alkyl, C₂ - C₆ alkenyl, C₁ - C₆ perfluoroalkyl, F, Cl, and Br, wherein:

10 R¹, R³, R⁴, R⁵ and R⁷ are independently selected from hydrogen and C₁ - C₆ alkyl;

R², R⁶, and R⁸ are independently selected from nitrogen protecting groups;

m and l are integers independently selected from 1 to 6; and

15 n is an integer from 0 to 2.

2. The method of claim 1 wherein Ar is phenyl substituted with said one or two groups.

3. The method of claim 1 wherein said nitrogen protecting group is -C(O)C₁-C₆ alkoxy.

20 4. The method of claim 1 wherein said nitrogen protecting group is benzylloxycarbonyl, fluorenyloxycarbonyl, acetyl, trifluoracetyl, chloroacetyl, benzoyl, t-butyloxycarbonyl, or benzyl.

5. The method of claim 1 wherein said compound of formula I is selected from the group consisting of

25 Methyl-(1-[4-[(4-oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-phenyl]-ethyl)-carbamic acid tert-butyl ester;

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[2-(2-Fluoro-4-[(4-oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-phenoxy)-ethyl]-propyl-carbamic acid tert-butyl ester;

Butyl-(2-{5-[(4-oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-pyridin-2-yloxy}-ethyl)-carbamic acid tert-butyl ester;

5 4-Oxo-4,5,6,7,8-hexahydro-cyclohepta[b]pyrrole-3-carboxylic acid (2-fluoro-4-hydroxy-phenyl)-amide;

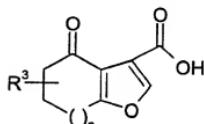
(1-Methyl-2-{4-[(4-oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-phenyl}-ethyl)-carbamic acid tert-butyl ester;

(2-{4-[(4-Oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-phenoxy}-ethyl)-propyl-carbamic acid tert-butyl ester; and

10 {2-Fluoro-5-[(4-oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-phenoxy}-acetic acid ethyl ester.

6. A method according to claim 1 further wherein said compound of formula II is prepared by

15 (a) reacting a compound of the formula



with an excess of an acid chloride or anhydride in a reaction inert solvent containing an excess of an acid acceptor until reaction is complete; and

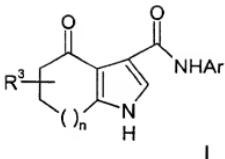
20 (b) adding an equivalent amount of NH₂-Ar to the solution of step (a) and holding until reaction is complete.

7. The method of claim 6 wherein said acid chloride is ethylchloroformate.

8. The method according to claim 1 which further comprises removing said nitrogen protecting group.

25 9. The method according to claim 5 which further comprises removing said nitrogen protecting group.

10. A compound of the following formula:



I

wherein Ar is phenyl or heterocycle, said phenyl or heterocycle being substituted with -O-(CH₂)_m-NR¹R², -O(CH₂)C(O)OR⁴, -CH(NR⁷R⁸)CH₃, -CH₂CH(NR⁵R⁶)CH₃, or OH, and said phenyl or heterocycle being optionally substituted with one or two groups selected from C₁ - C₆ alkoxy, C₁ - C₆ alkyl, C₂ - C₆ alkenyl, C₁ - C₆ perfluoroalkyl, F, Cl, and Br, wherein:

R¹, R³, R⁴, R⁵ and R⁷ are independently selected from hydrogen and C₁ - C₆ alkyl;

R², R⁶, and R⁸ are independently selected from nitrogen protecting groups;

m and l are integers independently selected from 1 to 6; and

n is an integer from 0 to 2.

10 11. A compound of claim 10 selected from the group consisting of:

Methyl-(1-[4-[(4-oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-phenyl]-ethyl)-carbamic acid tert-butyl ester;

[2-(2-Fluoro-4-[(4-oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-phenoxy)-ethyl]-propyl-carbamic acid tert-butyl ester;

Butyl-(2-[5-[(4-oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-pyridin-2-yloxy]-ethyl)-carbamic acid tert-butyl ester;

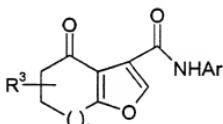
4-Oxo-4,5,6,7,8-hexahydro-cyclohepta[b]pyrrole-3-carboxylic acid (2-fluoro-4-hydroxy-phenyl)-amide;

(1-Methyl-2-{4-[(4-oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-phenyl}-ethyl)-carbamic acid tert-butyl ester;

(2-{4-[(4-Oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-phenoxy}-ethyl)-propyl-carbamic acid tert-butyl ester; and

{2-Fluoro-5-[(4-oxo-4,5,6,7-tetrahydro-1H-indole-3-carbonyl)-amino]-phenoxy}-acetic acid ethyl ester.

25 12. A compound of the following formula:



II

wherein Ar is phenyl or heterocycle, said phenyl or heterocycle being substituted with -O-(CH₂)_m-NR¹R², -O(CH₂)C(O)OR⁴, -CH(NR⁷R⁸)CH₃, -CH₂CH(NR⁵R⁶)CH₃, or OH, and said phenyl or heterocycle being optionally substituted with one or two groups selected from C₁ - C₆ alkoxy, C₁ - C₆ alkyl, C₂ - C₆ alkenyl, C₁ - C₆ perfluoroalkyl, F, Cl, and Br, wherein:

5 R¹, R³, R⁴, R⁵ and R⁷ are independently selected from hydrogen and C₁ - C₆ alkyl;

R², R⁶, and R⁸ are independently selected from nitrogen protecting groups;

m and l are integers independently selected from 1 to 6; and

n is an integer from 0 to 2.

13. The compound of claim 12 selected from the group consisting of:

10 Methyl-(1-[4-[(4-oxo-4,5,6,7-tetrahydro-benzofuran-3-carbonyl)-amino]-phenyl]-ethyl)-carbamic acid tert-butyl ester;

[2-(2-Fluoro-4-[(4-oxo-4,5,6,7-tetrahydro-benzofuran-3-carbonyl)-amino]-phenoxy)-ethyl]-propyl-carbamic acid tert-butyl ester;

15 Butyl-(2-(5-[(4-oxo-4,5,6,7-tetrahydro-benzofuran-3-carbonyl)-amino]-pyridin-2-yloxy)-ethyl)-carbamic acid tert-butyl ester;

4-Oxo-4,5,6,7,8-hexahydro-cyclohepta[b]furan-3-carboxylic acid (2-fluoro-4-hydroxy-phenyl)-amide;

(1-Methyl-2-[(4-oxo-4,5,6,7-tetrahydrobenzofuran-3-carbonyl)-amino]-phenyl)-ethyl)-carbamic acid tert-butyl ester;

20 (2-[(4-[(4-Oxo-4,5,6,7-tetrahydrobenzofuran-3-carbonyl)-amino]-phenoxy)-ethyl]-propyl-carbamic acid tert-butyl ester; and

(2-Fluoro-5-[(4-oxo-4,5,6,7-tetrahydrobenzofuran-3-carbonyl)-amino]-phenoxy)-acetic acid ethyl ester.

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